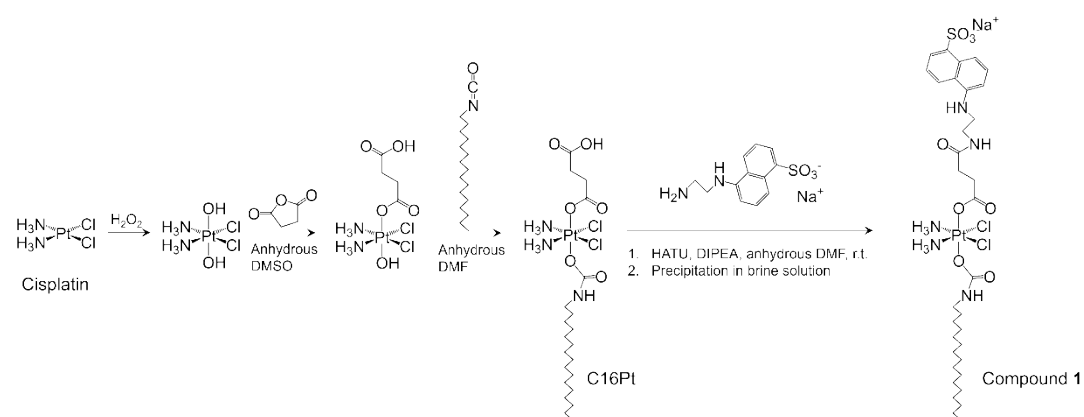


## Interactions between Mitochondria-Damaging Platinum(IV) Prodrugs and Cytochrome c

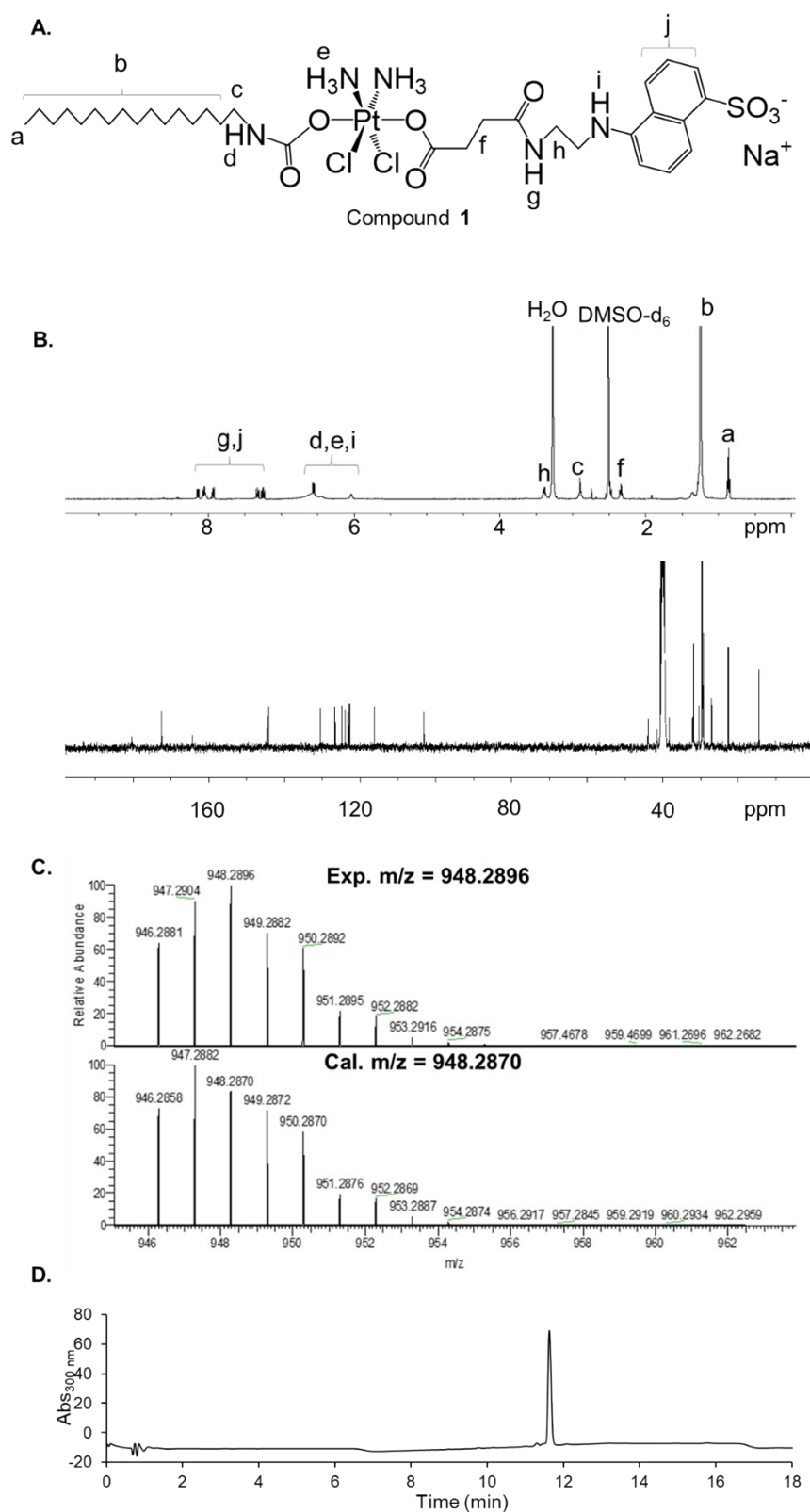
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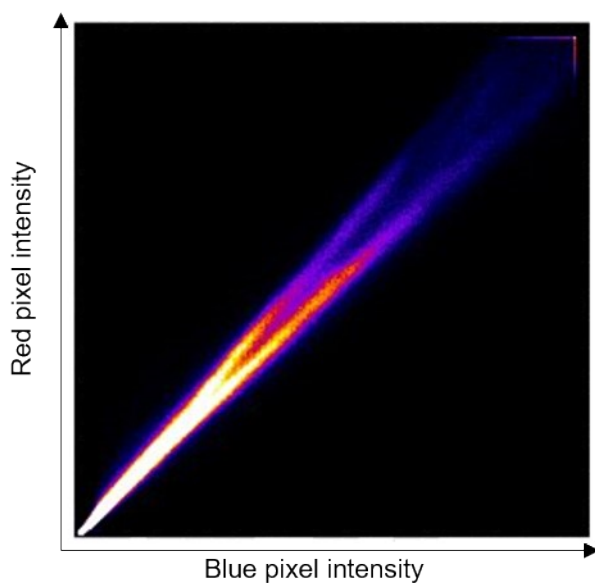
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**Fig S1.** Synthesis of the amphiphilic Pt(IV) prodrug (1).



**Fig S2.** Characterization of Compound 1: **A.**  $^1\text{H}$  NMR spectrum in  $\text{DMSO-d}_6$ ; **B.**  $^{13}\text{C}$  NMR spectrum in  $\text{DMSO-d}_6$ ; **C.** High resolution ESI-MS spectrum; **D.** Analytic HPLC analysis (Gradient: 0 min 5% B, 5 min 5% B, 10 min 95% B, 15 min 5% B. solvent A is 0.1% TFA aqueous solution and B is acetonitrile).



**Fig S3.** Colocalization analysis of **1** (blue) and Mitotracker Red (red) in HeLa cells. Pearson's correlation coefficient (PCC) = 0.993.

**Table S1:** Concentration values of cytochrome c (measured values are based on UV-vis absorbance data) and Compound **1** (measured with GFAAS) after dialyzed.

Sample	[Cyt c]/ $\mu\text{M}$ used	[ <b>1</b> ]/ $\mu\text{M}$ used	[Cyt c]/ $\mu\text{M}$ measured	[ <b>1</b> ]/ $\mu\text{M}$ measured	Ratio ([ <b>1</b> ]/[Cyt c])
Cyt c	50	0	50.0	0	0.0
Cyt c + 1 equiv. <b>1</b>	50	50	48.5	49.5	1.0
Cyt c + 2 equiv. <b>1</b>	50	100	48.1	97.8	2.0
Cyt c + 3 equiv. <b>1</b>	50	150	47.6	145.4	3.1
Cyt c + 4 equiv. <b>1</b>	50	200	47.1	195.2	4.1
Cyt c + 5 equiv. <b>1</b>	50	250	45.7	193.7	4.2